International application No.

|  | INTERNATIONAL SEARCH REPORT  | international application No. |   |  |  |  |  |
|--|--|-------------------------------|---|--|--|--|--|
|  |  |                               | PCT/JP2004/004696   |  |  |  |  |
| A. CLASSIFICATION OF SUBJECT MATTER Int.Cl <sup>7</sup> C07K16/28, C12N15/62, A61K39/395, A61P1/04, 1/16, 3/10, 5/14, 7/00, 7/04, 7/06, 11/06, 15/08, 17/00, 17/06, 21/04, 25/00, 27/02, 29/00, 35/00, 35/02, 37/02, 43/00 According to International Patent Classification (IPC) or to both national classification and IPC |  |                               |   |  |  |  |  |
| B. FIELDS SEA  |  |                               |   |  |  |  |  |
| Minimum documentation searched (classification system followed by classification symbols)  Int.Cl <sup>7</sup> C07K16/28, C12N15/62, A61K39/395, A61F1/04, 1/16, 3/10, 5/14,  7/00, 7/04, 7/06, 11/06, 15/08, 17/00, 17/06, 21/04, 25/00,  27/02, 29/00, 35/00, 35/02, 37/02, 43/00  |  |                               |   |  |  |  |  |
|  | earched other than minimum documentation to the exten  |                               |   |  | ·  |  |  |
| MEDLINE  | ase consulted during the international search (name of de<br>E(STN), BIOSIS(STN), WPIDS(STN),<br>cot/PIR/GeneSeq, Genbank/EMBL/DI                      | , JICS7                       | r file  | racticable, search   | n terms used)  |  |  |
| C. DOCUMENT  | TS CONSIDERED TO BE RELEVANT   |                               |   | ·  |  |  |  |
| Category*  | Citation of document, with indication, where app   | ropriate, of                  | f the relev   | ant passages   | Relevant to claim No.  |  |  |
| X<br>Y   | WO 01/97858 A2 (IDEC Pharmaceuticals Corp.), 1,2,4-13<br>27 December, 2001 (27.12.01), 3<br>& JP 2004-512262 A & EP 1299128 A2<br>& US 2002/0039557 A1 |                               |   |  |  |  |  |
| X<br>Y   | WO 02/22212 A2 (IDEC Pharmaceuticals Corp.), 1,2,4-13<br>21 March, 2002 (21.03.02),<br>& JP 2004-508420 A & EP 1328320 A2<br>& US 2002/0058029 A1      |                               |   | 1,2,4-13   |  |  |  |
| X  | WO 01/74388 Al (IDEC Pharmaceuticals Corp.), 11 October, 2001 (11.10.01), & JP 2004-500412 A & EP 1283722 Al & US 2002/0012665 Al                      |                               |   |  |  |  |  |
|  |  |                               |   |  |  |  |  |
| Further documents are listed in the continuation of Box C.   |  |                               |   |  |  |  |  |
| Special categories of cited documents:     "A" document defining the general state of the art which is not considered to be of particular relevance  |  | date                          | and not in  | oublished after the<br>conflict with the ap<br>theory underlying | international filing date or priority plication but cited to understand the invention                |  |  |
| "E" earlier application or patent but published on or after the international filing date  |  |                               | idered nov  | rticular relevance;<br>el or cannot be co<br>ocument is taken a  | the claimed invention cannot be<br>insidered to involve an inventive<br>lone                         |  |  |
| "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)  |  |                               | sidered to i  | involve an invent  | the claimed invention cannot be<br>ive step when the document is<br>such documents, such combination |  |  |
| "O" document referring to an oral disclosure, use, exhibition or other means document published prior to the international filing date but later than the priority date claimed  |  |                               | g obvious t   | o a person skilled in the same pate.                             | n the art<br>ent family  |  |  |
| Date of the actual completion of the international search 06 July, 2004 (06.07.04)   |  |                               | Date of mailing of the international search report 27 July, 2004 (27.07.04) |  |  |  |  |

Authorized officer

Telephone No.

Name and mailing address of the ISA/ Japanese Patent Office

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|                 |   | JP2004/004696           |
|-----------------|---|-------------------------|
| C (Continuation | ). DOCUMENTS CONSIDERED TO BE RELEVANT  |                         |
| Category*       | Citation of document, with indication, where appropriate, of the relevant passages  | Relevant to claim No.   |
| X<br>Y          | WO 02/04021 A1 (IDEC Pharmaceuticals Corp.),<br>17 January, 2002 (17.01.02),<br>& JP 2004-502742 A & EP 1305045 A1<br>& US 2002/0028178 A1  | 1,2,4-13                |
| X<br>Y          | JP 2001-518930 A (Immunomedics, Inc.),<br>16 October, 2001 (16.10.01),<br>& WO 98/42378 A1 & EP 969866 A1<br>& US 2002/0071807 A1           | 1,4,6-11<br>2,3,5,12,13 |
| X<br>Y          | JP 2002-544173 A (Immunomedics, Inc.),<br>24 December, 2002 (24.12.02),<br>& WO 00/67795 A1 & EP 1178826 A1                                 | 1,4,6-11<br>2,3,5,12,13 |
| X<br>Y          | JP 10-505231 A (Immunomedics, Inc.),<br>26 May, 1998 (26.05.98),<br>& WO 96/04925 A1 & EP 771208 A1<br>& US 5789554 A                       | 1,4,6-11<br>2,3,5,12,13 |
| Y               | HOLLIGER P. et al., "Diabodies": small bivalent and bispecific antibody fragments., Proc.Natl. Acad.Sci.USA., 1993, No.90, Vol.14, p.6444-8 | 2,3,5,12,13             |
| P,X             | WO 03/33654 A2 (ROSSI Edmund),<br>24 April, 2003 (24.04.03),<br>& US 2003/0148409 A1  | 1-13                    |
|                 |   |                         |
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| Box No. 1 Nucleotide and/or amino acid sequence(s) (Continuation of item1.b of the first sheet)  | 十   |
|--|-----|
| 1. With regard to any nucleotide and/or amino acid sequence disclosed in the international application and necessary to the claimed invention, the international search was carried out on the basis of:                           |     |
| a. type of material  |     |
| a sequence listing   |     |
| table(s) related to the sequence listing   |     |
| b. format of material  |     |
| in written format  |     |
| in computer readable form  |     |
| c. time of filing/furnishing   |     |
| contained in the international application as filed  |     |
| illed together with the international application in computer readable form  |     |
| furnished subsequently to this Authority for the purposes of search  |     |
| 2. X In addition, in the case that more than one version or copy of a sequence listing and/or table relating thereto has been filed  | 1   |
| or furnished, the required statements that the information in the subsequent or additional copies is identical to that in the application as filed or does not go beyond the application as filed, as appropriate, were furnished. |     |
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| 3. Additional comments:  | 1   |
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| Box No. 1   | Observations where certain claims were found unsearchable (Continuation of item 2 of first sheet)  |  |  |  |  |
|---|--|--|--|--|--|
| This international search report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:  |  |  |  |  |  |
|   | Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:  |  |  |  |  |
|   |  |  |  |  |  |
| 1   | Claims Nos.: because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically: |  |  |  |  |
| ·,  |  |  |  |  |  |
|   | Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).   |  |  |  |  |
| Box No. 1   | Observations where unity of invention is lacking (Continuation of item 3 of first sheet)   |  |  |  |  |
| This International Searching Authority found multiple inventions in this international application, as follows:  It is recognized that degraded antibodies recognizing CD22, i.e., the matter common to claims 1 to 13, have been publicly known (see, if needed, WO98/42378). Thus, the above common matter cannot be considered as a special technical feature. Such being the case, the inventions as set forth in claims 1 to 13 cannot be regarded as a group so linked as to form a single general inventive concept.  Thus, the inventions as set forth in claims 1 to 13 are classified into the following 4 groups of inventions: (1) a degraded antibody having the amino acid sequence represented by SEQ ID NO:1, (2) a degraded antibody having the amino acid sequence represented by SEQ ID NO:3, (continued to extra sheet) |  |  |  |  |  |
| searchabl   | As all required additional search fees were timely paid by the applicant, this international search report covers all e claims.  |  |  |  |  |
|   | As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.   |  |  |  |  |
| 3. Covers   | As only some of the required additional search fees were timely paid by the applicant, this international search report  |  |  |  |  |
|   | only those claims for which fees were paid, specifically claims Nos.:  |  |  |  |  |
| 4. No required additional search fees were timely paid by the applicant. Consequently, this international search report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:  The parts relating to SEQ ID NO:1 in claims 1 to 13  |  |  |  |  |  |
| Remark (  | on Protest  The additional search fees were accompanied by the applicant's protest.  No protest accompanied the payment of additional search fees.   |  |  |  |  |

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| Continuation of   | f Box No.III of cont  | inuation of first s   | heet(2)   |
|---|---|---|---|
| (3) a degraded ant<br>sequence represen<br>having CDR of SEQ<br>SEQ ID NO:11. | tibody having CDR of<br>ted by SEQ ID NO:<br>ID NO:9 and the am | f SEQ ID NO:5 and to<br>7, and (4) a degra<br>ino acid sequence r | he amino acid<br>aded antibody<br>epresented by |
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Form PCT/ISA/210 (extra sheet) (January 2004)